Applicant: Kornelia Polyak et al.

Attorney's Docket No.: 00530-094001 / DFCI 689 / MGH 1897

Serial No.: 10/081,817 Filed: February 22, 2002

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## Amendments to the Claims:

This listing replaces all prior versions and listings of claims in the application:

## 1-22. (Cancelled)

23. (Currently amended) A method of diagnosis, the method comprising:

(a) providing a test cell; and

(b) determining, in the test cell, the degree of methylation of a HIN-1 promoter region in the test cell one or more C residues in a nucleotide sequence consisting of (i) the HIN-1 promoter region and (ii) the first twelve nucleotides of SEQ ID NO:3,

wherein the C residues are C residues in CpG sequences, and

wherein a high degree of methylation of the HIN 1 promoter region <u>C</u> residues is an indication that the test cell is a cancer cell.

- 24. (Original) The method of claim 23, wherein the test cell is a breast cell.
- 25. (Withdrawn) An isolated polypeptide comprising (a) a functional fragment of the polypeptide of claim 8; or (b) the functional fragment, except for one or more conservative amino acid substitutions.
- 26. (Withdrawn) An isolated DNA comprising a fragment of the nucleic acid with SEQ ID NO:3, wherein the fragment comprises nucleotides 55 and 56 of SEQ ID NO:3.
  - 27. (Withdrawn) An antibody that binds to the polypeptide of claim 8.
- 28. (Withdrawn) The antibody of claim 27, wherein the antibody is a monoclonal antibody.
- 29. (Withdrawn) The antibody of claim 27, wherein the antibody is a polyclonal antibody.

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30. (Withdrawn) A method of treatment comprising identifying a patient as having cancer cells in which (a) HIN-1 gene expression is low or (b) a HIN-1 promoter region is methylated; and treating the patient with a compound that reduces methylation of the HIN-1 promoter region.

- 31. (Withdrawn) A method of identifying a compound that replaces the function of HIN-1 in cells that do not express HIN-1, the method comprising:
  - (a) providing a first cell that does not express HIN-1;
  - (b) providing a second cell that does express HIN-1;
  - (c) treating the first cell and the second cell with a test compound; and
- (d) determining whether the test compound decreases proliferation of the first or the second cell, wherein a compound that decreases proliferation of the first cell but not the second cell can potentially replace the function of HIN-1 in cells that do not express HIN -1.
  - 32. (Withdrawn) A method of treatment comprising

identifying a patient as having cancer cells in which (a) HIN-1 gene expression is low or (b) a HIN-1 promoter region is methylated; and

treating the patient with a compound that induces expression of a gene with a methylated promoter region.

- 33. (Withdrawn) The method of claim 23, wherein the cell is a pancreatic cell.
- 34. (Withdrawn) The method of claim 23, wherein the cell is a prostate cell.
- 35. (New) The method of claim 23, wherein the test cell is selected from the group consisting of a lung cell, a prostate cell, a pancreatic cell, a gastrointestinal cell, and a skin cell.
- 36. (New) The method of claim 23, wherein the promoter region comprises SEQ ID NO:19.
- 37 (New) The method of claim 36, wherein the promoter region consists of SEQ ID NO:19.

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- 38. (New) The method of claim 23, wherein the segment comprises nucleotide 1 to nucleotide 252 of SEQ ID NO:19.
- 39. (New) The method of claim 23, wherein the segment consists of nucleotide 1 to nucleotide 252 of SEQ ID NO:19.
- 40. (New) The method of claim 23, wherein the segment comprises nucleotide 229 to nucleotide 551 of SEQ ID NO:19 and nucleotide 1 to nucleotide 12 of SEQ ID NO:3.
- 41. (New) The method of claim 23, wherein the segment consists of nucleotide 229 to nucleotide 551 of SEQ ID NO:19 and nucleotide 1 to nucleotide 12 of SEQ ID NO:3.
- 42. (New) The method of claim 23, wherein the segment comprises SEQ ID NO:19 and nucleotide 1 to nucleotide 12 of SEQ ID NO:3.
- 43. (New) The method of claim 23, wherein the segment consists of SEQ ID NO:19 and nucleotide 1 to nucleotide 12 of SEQ ID NO:3.
  - 44. (New) The method of claim 23, wherein the test cell is a human cell.
- 45. (New) The method of claim 23, wherein the degree of methylation is determined by sequencing of bisulfite-treated DNA comprising the segment.
- 46. (New) The method of claim 23, wherein the degree of methylation is determined by a methylation-specific polymerase chain reaction (MCP) assay.